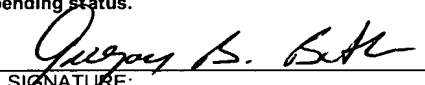


<b>TRANSMITTAL LETTER TO THE UNITED STATES DESIGNATED/ELECTED OFFICE (DO/EO/US) CONCERNING A SUBMISSION UNDER 35 U.S.C. 371</b>		ATTORNEY'S DOCKET NUMBER 63628 (46342)
INTERNATIONAL APPLICATION NO. PCT/JP2003/016978		INTERNATIONAL FILING DATE 26 December 2003
TITLE OF INVENTION METASTIN DERIVATIVES AND USE THEREOF		PRIORITY DATE CLAIMED 26 December 2002
APPLICANT(S) FOR DO/EO/US Chieko Kitada et al.		
Applicant herewith submits to the United States Designated/Elected Office (DO/EO/US) the following items and other information:		
<ol style="list-style-type: none"> <li>1. <input checked="" type="checkbox"/> This is a <b>FIRST</b> submission of items concerning a submission under 35 U.S.C. 371.</li> <li>2. <input type="checkbox"/> This is a <b>SECOND</b> or <b>SUBSEQUENT</b> submission of items concerning a submission under 35 U.S.C. 371.</li> <li>3. <input type="checkbox"/> This is an express request to begin national examination procedures (35 U.S.C. 371 (f)). The submission must include items (5), (6), (9) and (21) indicated below.</li> <li>4. <input type="checkbox"/> The US has been elected (Article 31).</li> <li>5. <input checked="" type="checkbox"/> A copy of the Cover Page of the International Application as filed (35 U.S.C. 371 (c)(2))               <ol style="list-style-type: none"> <li>a. <input checked="" type="checkbox"/> is attached hereto (required only if not communicated by the International Bureau).</li> <li>b. <input type="checkbox"/> has been communicated by the International Bureau.</li> <li>c. <input type="checkbox"/> is not required, as the application was filed in the United States Receiving Office (RO/US).</li> </ol> </li> <li>6. <input checked="" type="checkbox"/> An English language translation of the International Application as filed (35 U.S.C. 371 (c)(2)).               <ol style="list-style-type: none"> <li>a. <input checked="" type="checkbox"/> is attached hereto.</li> <li>b. <input type="checkbox"/> has been previously submitted under 35 U.S.C. 154(d)(4).</li> </ol> </li> <li>7. <input checked="" type="checkbox"/> Amendments to the claims of the International Application under PCT Article 19 (35 U.S.C. 371 (c)(3))               <ol style="list-style-type: none"> <li>a. <input checked="" type="checkbox"/> are attached hereto (required only if not communicated by the International Bureau).</li> <li>b. <input type="checkbox"/> have been communicated by the International Bureau.</li> <li>c. <input type="checkbox"/> have not been made; however, the time limit for making such amendments has NOT expired.</li> <li>d. <input type="checkbox"/> have not been made and will not be made.</li> </ol> </li> <li>8. <input type="checkbox"/> An English language translation of the amendments to the claims under PCT Article 19 (35 U.S.C. 371 (c)(3)).</li> <li>9. <input type="checkbox"/> An oath or declaration of the inventor(s) (35 U.S.C. 371 (c)(4)).</li> <li>10. <input type="checkbox"/> An English language translation of the annexes of the International Preliminary Examination Report under PCT Article 36 (35 U.S.C. 371 (c)(5)).</li> </ol>		
<b>Items 11 to 20 below concern document(s) or information included:</b>		
<ol style="list-style-type: none"> <li>11. <input checked="" type="checkbox"/> An Information Disclosure Statement under 37 CFR 1.97 and 1.98.</li> <li>12. <input type="checkbox"/> An assignment document for recording. A separate cover sheet in compliance with 37 CFR 3.28 and 3.31 is included.</li> <li>13. <input checked="" type="checkbox"/> A preliminary amendment.</li> <li>14. <input type="checkbox"/> An Application Data Sheet under 37 CFR 1.76.</li> <li>15. <input type="checkbox"/> A substitute specification.</li> <li>16. <input type="checkbox"/> A power of attorney and/or change of address letter.</li> <li>17. <input checked="" type="checkbox"/> A computer-readable form of the sequence listing in accordance with PCT Rule 13ter.2 and 37 CFR 1.821 - 1.825.</li> <li>18. <input type="checkbox"/> A second copy of the published International Application under 35 U.S.C. 154(d)(4).</li> <li>19. <input type="checkbox"/> A second copy of the English language translation of the international application under 35 U.S.C. 154(d)(4).</li> <li>20. <input checked="" type="checkbox"/> Other items or information: PCT/RO/101; PCT/IB/304; PCT/IB/306; PCT/ISA/210; PCT/IPEA/409; PCT/IPEA/416; Receipt in the Case of an Original Deposit (2 documents); Sequence Listing (14 pages); Return Receipt Postcard</li> </ol>		

Under the Paperwork Reduction Act of 1995, no persons are required to respond to a collection of information unless it displays a valid OMB control number.

U.S. APPLICATION NO. <b>10/540494</b>		INTERNATIONAL APPLICATION NO. PCT/JP2003/016978		ATTORNEY'S DOCKET NUMBER 63628 (46342)	
The following fees have been submitted				CALCULATIONS	PTO USE ONLY
21.	<input checked="" type="checkbox"/>	Basic national fee .....	\$300	\$ 300.00	
22.	<input checked="" type="checkbox"/>	Examination fee			
If International preliminary examination report prepared by USPTO and all claims satisfy provisions of PCT Article 33(1)-(4) .....				\$100	
All other situations .....				\$200	
				\$ 200.00	
23.	<input checked="" type="checkbox"/>	Search fee			
Search fee (37 CFR 1.445(a)(2)) has been paid on the international application to the USPTO as an International Searching Authority .....				\$100	
International Search Report prepared and provided to the Office .....				\$400	
All other situations .....				\$500	
TOTAL OF 21, 22 and 23 =				\$ 1,000.00	
<input type="checkbox"/> Additional fee for specification and drawings filed in paper over 100 sheets (excluding sequence listing or computer program listing filed in an electronic medium). The fee is \$250 for each additional 50 sheets or fraction thereof.					
Total Sheets	Extra sheets	Number of each additional 50 or fraction thereof (round up to a whole number)	RATE		
- 100 =	/50 =		x \$250.00		
Surcharge of \$130.00 for furnishing the oath or declaration later than 30 months from the earliest claimed priority date (37 CFR 1.492 (e)).				\$ 130.00	
CLAIMS	NUMBER FILED	NUMBER EXTRA	RATE		
Total claims	47 - 20 =	27	x 50.00	\$ 1,350.00	
Independent claims	11 - 3 =	8	x 200.00	\$ 1,600.00	
MULTIPLE DEPENDENT CLAIM(s) (if applicable)			+ 360.00	\$ 360.00	
TOTAL OF ABOVE CALCULATIONS =				\$ 4,440.00	
<input type="checkbox"/> Applicant claims small entity status. See 37 CFR 1.27. Fees above are reduced by 1/2.				\$	
SUBTOTAL =				\$ 4,440.00	
Processing fee of \$130.00 for furnishing the English translation later than 30 months from the earliest claimed priority date (37 CFR 1.492 (f)).				\$	
TOTAL NATIONAL FEE =				\$ 4,440.00	
Fee for recording the enclosed assignment (37 CFR 1.21 (h)). The assignment must be accompanied by an appropriate cover sheet (37 CFR 3.28, 3.31). \$40.00 per property				\$	
TOTAL FEES ENCLOSED =				\$	
				Amount to be refunded:	\$
				Amount to be charged:	\$4,440.00
a. <input type="checkbox"/> A check in the amount of \$ _____ to cover the above fees is enclosed. b. <input checked="" type="checkbox"/> Please charge my Deposit Account No. <u>04-1105</u> in the amount of \$ <u>4,440.00</u> to cover the above fees. A duplicate copy of this sheet is enclosed. c. <input checked="" type="checkbox"/> The Commissioner is hereby authorized to charge any additional fees which may be required, or credit any overpayment to Deposit Account No. <u>04-1105</u> . A duplicate copy of this sheet is enclosed. d. <input type="checkbox"/> Fees are to be charged to a credit card. <b>WARNING:</b> Information on this form may become public. <b>Credit card information should not be included on this form.</b> Provide credit card information and authorization on PTO-2038.					
<b>NOTE:</b> Where an appropriate time limit under 37 CFR 1.495 has not been met, a petition to revive (37 CFR 1.137 (a) or (b)) must be filed and granted to restore the International Application to pending status.					
SEND ALL CORRESPONDENCE TO:					
David G. Conlin EDWARDS & ANGELL, LLP P.O. Box 55874 Boston, Massachusetts 02205 (617) 439-4444			SIGNATURE:  Gregory B. Butler, Ph.D. NAME 34,558 REGISTRATION NUMBER		
CUSTOMER NUMBER: 21874					

Application No. (if known): Not Yet Assigned

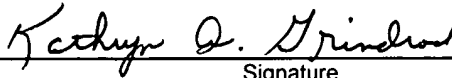
Attorney Docket No.: 63628 (46342)

**Certificate of Express Mailing Under 37 CFR 1.10**

I hereby certify that this correspondence is being deposited with the United States Postal Service as Express Mail, Airbill No. EV 654381958 US in an envelope addressed to:

MS PCT  
Commissioner for Patents  
P.O. Box 1450  
Alexandria, VA 22313-1450

on June 23, 2005  
Date



Signature

Kathryn A. Grindrod

Typed or printed name of person signing Certificate

Registration Number, if applicable

(617) 517-5534

Telephone Number

Note: Each paper must have its own certificate of mailing, or this certificate must identify each submitted paper.

Transmittal Letter to the United States Designated-Elected Office (2 pages)  
Specification Cover Sheet (1 page)  
Specification excluding claims (233 pages)  
47 claims (10 pages)  
Abstract (3 pages)  
Preliminary Amendment ( pages)  
Formal Drawings Fig. 1 - Fig.22 (22 sheets)  
PCT/RO/101; PCT/IB/304; PCT/IB/306; PCT/ISA/210; PCT/IPEA/409;  
PCT/IPEA/416; Receipt in the Case of an Original Deposit (2 documents)  
Sequence Statement (2 pages)  
Sequence Listing (14 pages)  
Computer readable disk containing Sequence Listing  
Information Disclosure Statement (2 pages)  
IDS (Citation) by Applicant (14 References) (1 page)  
This Return Receipt Postcard  
Charge \$4,440.00 to deposit account 04-1105